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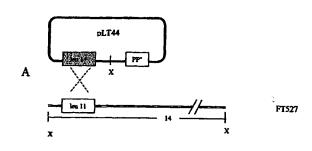
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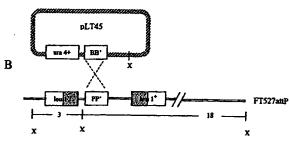
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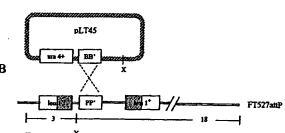
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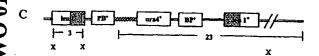
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(54) Title: DNA RECOMBINATION IN EUKARYOTIC CELLS BY THE BACTERIOPHAGE PHIC31 RECOMBINATION SYS-TEM









(57) Abstract: This invention provides methods for obtaining specific and stable integration of nucleic acids into eukaryotic cells. The invention makes use of site-specific recombination systems that use prokaryotic recombinase polypeptides, such as the ΦC31 integrase, that can mediate recombination between the recombination sites, but not between hybrid recombination sites that are formed upon the recombination. Thus, the recombination is irreversible in the absence of additional factors. Eukaryotic cells that contain the recombinase polypeptides, or genes that encode the recombinases, are also provided.



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A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12N5/10 C12N C12N15/79 C12N15/90 //C12N9/22 According to International Patent Classification (IPC) or to both national classification and IPC **B. FIELDS SEARCHED** Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12N Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, WPI Data, PAJ, BIOSIS, CHEM ABS Data, EMBASE C. DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. P,X WO OO 11155 A (UNIV LELAND STANFORD 1-8 JUNIOR) 2 March 2000 (2000-03-02) 10-47 page 4 page 5, line 25 -page 6, line 2 A THORPE, H. M. ET AL.: "In vitro 1-47 site-specific integration of bacteriophage DNA catalyzed by a recombinase of the resolvase/invertase family" vol. 95, 1998, pages 5505-5510, XP002157128 cited in the application abstract -/--Further documents are listed in the continuation of box C. Patent family members are listed in annex. Special categories of cited documents: \*T\* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the hearth. "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone \*L\* document which may throw doubts on priority claim(s) or which is clied to establish the publication date of another citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled "O" document referring to an oral disclosure, use, exhibition or document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 12 January 2001 29/01/2001 Name and mailing address of the ISA Authorized officer European Patenti Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Filijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nil, Fac: (+31-70) 340-3016 Mata Vicente, T.

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| Category * | Citation of document, with indication, where appropriate, of the relevant passages   | Relevant to claim No. |
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| 4          | WO 99 18222 A (ALONSO JUAN C ; ROJO<br>FERNANDO (ES); BERNARD ANTONIO (ES); DIAZ<br>VICE) 15 April 1999 (1999-04-15)<br>page 14, paragraph 2 | 1-47                  |
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## INTERNATIONAL SEARCH REPORT

information on patent family members

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| Patent document<br>cited in search report |   | Publication date | Patent family member(s) |                        | Publication date         |
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